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June 17, 2003

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Commissioner for Patents  
P.O. Box 1450  
Alexandria, VA 22313-1450

**Art Unit 1634**

Re: U.S. Utility Patent Application  
Appl. No. 09/695,065; Filed: October 25, 2000  
For: **Methods of Manipulating and Sequencing Nucleic Acid Molecules Using Transposition and Recombination**  
Inventors: BRASCH *et al.*  
Our Ref: 0942.5000001/RWE/BJD/JKM

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Sir:

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Transmitted herewith for appropriate action are the following documents:

1. Second Supplemental Information Disclosure Statement;
2. List of cited references on Form PTO-1449 (21 sheets);
3. A copy of each reference cited on Form PTO-1449 (76 documents: **AM6-AL9, and AR85-AR105**); and
4. One (1) return postcard.

It is respectfully requested that the attached postcard be stamped with the date of filing of these documents, and that it be returned to our courier. In the event that extensions of time are necessary to prevent abandonment of this patent application, then such extensions of time are hereby petitioned.

Commissioner for Patents  
June 17, 2003  
Page 2

The U.S. Patent and Trademark Office is hereby authorized to charge any fee deficiency, or credit any overpayment, to our Deposit Account No. 19-0036.

Respectfully submitted,

STERNE, KESSLER, GOLDSTEIN & FOX P.L.L.C.

A handwritten signature in black ink, appearing to read "Brian J. Del Buono", written over a horizontal line.

Brian J. Del Buono  
Attorney for Applicants  
Registration No. 42,473

RWE/BJD/JKM/acr  
Enclosures

SKGF\_DC1:138517.1



IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re application of:

Brasch *et al.*

Appl. No. 09/695,065

Filed: October 25, 2000

For: **Methods of Manipulating and  
Sequencing Nucleic Acid  
Molecules Using Transposition  
and Recombination**

Confirmation No. 1682

Art Unit: 1634

Examiner: Lu, F.

Atty. Docket: 0942.5000001/RWE/BJD/JKM

### Second Supplemental Information Disclosure Statement

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Sir:

Listed on accompanying Form PTO-1449 are documents that may be considered material to the examination of this application, in compliance with the duty of disclosure requirements of 37 C.F.R. §§ 1.56, 1.97 and 1.98. The numbering on this Second Supplemental Information Disclosure Statement is a continuation of the numbering in Applicants' Information Disclosure Statement filed on June 12, 2001, and First Supplemental Information Disclosure Statement filed electronically herewith, in connection with the above-captioned application. A copy of each document is also provided.

Where the publication date of a listed document does not provide a month of publication, the year of publication of the listed document is sufficiently earlier than the effective U.S. filing date and any foreign priority date so that the month of publication is not in issue. Applicants have listed publication dates on the attached PTO-1449 based on information presently available to the undersigned. However, the listed publication dates

should not be construed as an admission that the information was actually published on the date indicated.

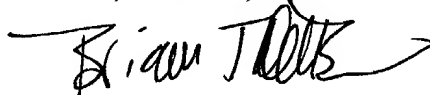
This Second Supplemental Information Disclosure Statement is being filed before the mailing of a first Office Action after the filing of a request for continued examination under 37 C.F.R. § 1.114. No statement or fee is required.

Applicants reserve the right to establish the patentability of the claimed invention over any of the information provided herewith, and/or to prove that this information may not be prior art, and/or to prove that this information may not be enabling for the teachings purportedly offered. This statement should not be construed as a representation that a search has been made, or that information more material to the examination of the present patent application does not exist.

Consideration of the cited documents and making the same of record in the prosecution of the above-identified application is respectfully requested. The U.S. Patent and Trademark Office is hereby authorized to charge any fee deficiency, or credit any overpayment, to our Deposit Account No. 19-0036.

Respectfully submitted,

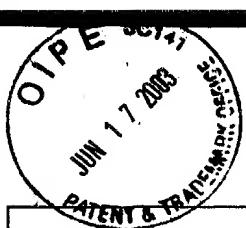
STERNE, KESSLER, GOLDSTEIN & FOX P.L.L.C.



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Date: June 17, 2003

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SECOND SUPPLEMENTAL  
INFORMATION DISCLOSURE STATEMENTATTY. DOCKET NO.  
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Brasch et al.FILING DATE  
October 25, 2000GROUP  
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## U.S. PATENT DOCUMENTS

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## FOREIGN PATENT DOCUMENTS

EXAMINER INITIAL		DOCUMENT NUMBER	DATE	COUNTRY	CLASS	SUB-CLASS	TRANSLATION
	AL						Yes No
	AM6	EP 0 220 009	04/29/1987	Europe			Yes No
	AN6	WO 90/11375	10/04/1990	WIPO			Yes No
	AO6	EP 0 427 074	05/15/1991	Europe			Yes No
	AP6	WO 91/09957	07/11/1991	WIPO			Yes No

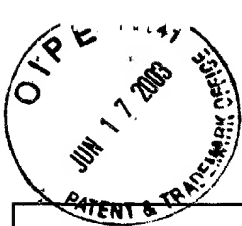
## OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	<u>85</u>	Akagi, K., et al., "Cre-mediated somatic site-specific recombination in mice," <i>Nucl. Acids Res.</i> 25:1781-1788, Oxford University Press (1997).
	AS	<u>85</u>	Aladjem, M.I., et al., "Positive Selection of FLP-Mediated Unequal Sister Chromatid Exchange Products in Mammalian Cells," <i>Mol. Cell. Biol.</i> 17:857-861, American Society for Microbiology (1997).
	AT	<u>85</u>	Angrand, P.-O., et al., "Inducible expression based on regulated recombination: a single vector strategy for stable expression in cultured cells," <i>Nucl. Acids Res.</i> 26:3263-3269, Oxford University Press (July 1998).

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	APPLICANT Brasch et al.	
	FILING DATE October 25, 2000	GROUP 1682

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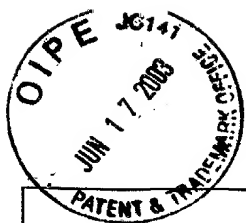
FOREIGN PATENT DOCUMENTS							
EXAMINER INITIAL		DOCUMENT NUMBER	DATE	COUNTRY	CLASS	SUB-CLASS	TRANSLATION
	AL7	WO 92/15694	09/17/1992	WIPO			Yes No
	AM7	WO 92/22650	12/23/1992	WIPO			Yes No
	AN7	EP 0 542 466	05/19/1993	Europe			Yes No
	AO7	WO 94/03624	02/17/1994	WIPO			Yes No
	AP7	WO 94/18333	08/18/1994	WIPO			Yes No

## OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	86	Ayres, E.K., et al., "Precise Deletions in Large Bacterial Genomes by Vector-mediated Excision (VEX). The trfA Gene of Promiscuous Plasmid RK2 is Essential for Replication in Several Gram-negative Hosts," <i>J. Mol. Biol.</i> 230:174-185, Academic Press Limited (1993).
	AS	86	Backman, K., et al., "Use of Synchronous Site-Specific Recombination In Vivo to Regulate Gene Expression," <i>Bio/Technology</i> 2:1045-1049, Nature Publishing Company (1984).
	AT	86	Bai, C., et al., "SKP1 Connects Cell Cycle Regulators to the Ubiquitin Proteolysis Machinery through a Novel Motif, the F-Box," <i>Cell</i> 86:263-274, Cell Press (1996).

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	AL8	WO 96/04393	02/15/1996	WIPO			Yes No
	AM8	WO 96/30498	10/03/1996	WIPO			Yes No
	AN8	WO 96/40724	12/19/1996	WIPO			Yes No
	AO8	WO 01/07572	02/01/2001	WIPO			Yes No
	AP8	WO 01/62892	08/30/2001	WIPO			Yes No

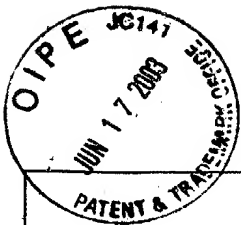
## OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	<u>87</u>	Bernard, P. et al., "The F Plasmid CcdB Protein Induces Efficient ATP-dependent DNA Cleavage by Gyrase," <i>J. Mol. Biol.</i> 234:534-541, Academic Press (1993).
	AS	<u>87</u>	Boshart, M., et al., "A Very Strong Enhancer is Located Upstream of an Immediate Early Gene of Human Cytomegalovirus," <i>Cell</i> 41:521-530, The MIT Press (1985).
	AT	<u>87</u>	Bouhassira, E.E., et al., "Transcriptional Behavior of LCR Enhancer Elements Integrated at the Same Chromosomal Locus by Recombinase-Mediated Cassette Exchange," <i>Blood</i> 90:3332-3344, The American Society of Hematology (1997).

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	AL9	WO 02/05294	01/17/2002	WIPO			Yes No
	AM						Yes No
	AN						Yes No
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	AP						Yes No

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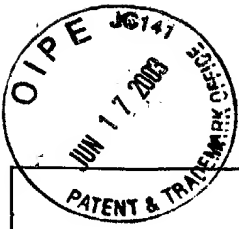
	AR	<u>88</u>	Choulika, A., et al., "Transfer of Single Gene-Containing Long Terminal Repeats into the Genome of Mammalian Cells by a Retroviral Vector Carrying the cre Gene and the loxP Site." <i>J. Virol.</i> 70:1792-1798, American Society for Microbiology (1996).
	AS	<u>88</u>	Cigan, A.M., et al., "Mutational Analysis of the HIS4 Translational Initiator Region in <i>Saccharomyces cerevisiae</i> ," <i>Mol. Cell. Biol.</i> 8:2964-2975, American Society for Microbiology (1988).
	AT	<u>88</u>	CLONTECH, "Creator™ Gene Cloning & Expression System," <i>CLONTECHniques</i> 15:7-11, CLONTECH (April 2000).

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	AL						Yes No
	AM						Yes No
	AN						Yes No
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## OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	<u>89</u>	CLONTECH, "New Additions to the Creator™ Platform," CLONTECHniques 16:4 pages, CLONTECH (January 2001).
	AS	<u>89</u>	CLONTECH, "New Creator™ -Compatible Expression Systems," CLONTECHniques 15:2 pages, CLONTECH (October 2000).
	AT	<u>89</u>	CLONTECH, "Creator™ Acceptor Vector Construction Kit," CLONTECHniques 16:2 pages, CLONTECH (October 2001).

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## OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	<u>90</u>	CLONTECH, "Creator™ SMART™ Library Construction Kit," CLONTECHniques 16:2 pages, CLONTECH (October 2001).
	AS	<u>90</u>	CLONTECH, "Creator™: The Universal Platform for Analysis of Gene Function," Powerpoint Presentation, 9 pages, CLONTECH, (July 24, 2001), available at <a href="http://www.clontech.com/products/families/creator/popups/slpage1.html">http://www.clontech.com/products/families/creator/popups/slpage1.html</a> .
	AT	<u>90</u>	CLONTECH, "Creator™ pDNR-Dual Cloning Kit," CLONTECHniques 16:3 pages, CLONTECH, (October 2001).

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## OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

	AR	<u>91</u>	Curcio, M.J., and Garfinkel, D.J., "Single-step selection for Ty1 element retrotransposition," <i>Proc. Natl. Acad. Sci. USA</i> 88:936-940, National Academy of Sciences (1991).
	AS	<u>91</u>	Dijkema, R., et al., "Cloning and expression of the chromosomal immune interferon gene of the rat," <i>EMBO J.</i> 4:761-767, IRL Press Limited (1985).
	AT	<u>91</u>	Flanagan, P.M. and Fennwald, M.A., "Analysis of Inhibitors of the Site-specific Recombination Reaction Mediated by Tn3 Resolvase," <i>J. Mol. Biol.</i> 206:295-304, Academic Press Limited (1989).

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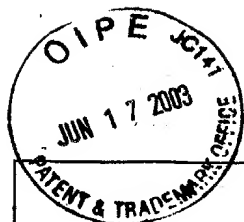
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	AR	92	Gay, P., et al., "Positive Selection Procedure for Entrapment of Insertion Sequence Elements in Gram-Negative Bacteria," <i>J. Bacteriol.</i> 164:918-921, American Society for Microbiology (1985).
	AS	92	Gay, P., et al., "Cloning Structural Gene <i>sacB</i> , Which Codes for Exoenzyme Levansucrase of <i>Bacillus subtilis</i> : Expression of the Gene in <i>Escherichia coli</i> ," <i>J. Bacteriol.</i> 153:1424-1431, American Society for Microbiology (1983).
	AT	92	Gorman, C.M., et al. "The Rous sarcoma virus long terminal repeat is a strong promoter when introduced into a variety of eukaryotic cells by DNA-mediated transfection," <i>Proc. Natl. Acad. Sci. USA</i> 79:6777-6781, National Academy of Sciences (1982).

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	AR	93	Grindly, N.D.F., and Kelley, W.S., "Effects of Different Alleles of the <i>E. coli</i> K12 <i>polA</i> Gene on the Replication of Non-transferring Plasmids," <i>Molec. Gen. Genet.</i> 143:311-318, Springer Verlag (1976).
	AS	93	Gronostajski, R.M., and Sadowski, P.D., "The FLP Protein of the 2-micron Plasmid of Yeast. Inter- and Intramolecular Reactions," <i>J. Biol. Chem.</i> 260:12328-12335, The American Society of Biological Chemists, Inc. (1985).
	AT	93	Haffter, P., and Bickle, T.A., "Enhancer-independent mutants of the <i>Cin</i> recombinase have a relaxed topological specificity," <i>EMBO J.</i> 7:3991-3996, IRL Press Limited (1988).

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AR	94	Hehl, R., et al., "Structural analysis of Tam3, a transposable element from <i>Antirrhinum majus</i> , reveals homologies to the Ac element from maize," <i>Plant Molec. Biol.</i> 16:369-371, Kluwer Academic Publishers (1991).
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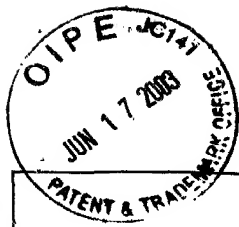
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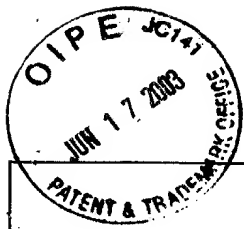
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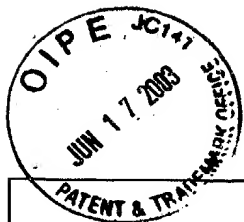
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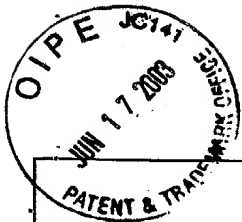
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	AR	<u>99</u>	Metcalfe, W.W., et al., "Conditionally Replicative and Conjugative Plasmids Carrying lacZ $\alpha$ for Cloning, Mutagenesis, and Allele Replacement in Bacteria," <i>Plasmid</i> 35:1-13, Academic Press (1996).
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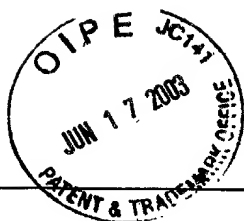
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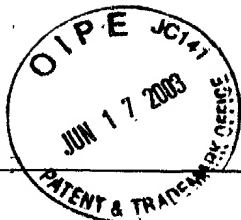
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	AR	<u>101</u>	Qin, M., et al., "Cre recombinase-mediated site-specific recombination between plant chromosomes," <i>Proc. Natl. Acad. Sci. USA</i> 91:1706-1710, National Academy of Sciences (1994).
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	AR	<u>102</u>	Sato, T., et al., "The <i>cisA</i> Cistron of <i>Bacillus subtilis</i> Sporulation Gene <i>spoIVC</i> Encodes a Protein Homologous to a Site-Specific Recombinase," <i>J. Bacteriol.</i> 172:1092-1098, American Society for Microbiology (1990).
	AS	<u>102</u>	Sauer, B., et al., "Construction of Isogenic Cell Lines Expressing Human and Rat Angiotensin II AT <sub>1</sub> Receptors by Cre-Mediated Site-Specific Recombination," <i>Methods: A Companion to Methods in Enzymology</i> 4:143-149, Academic Press (1992).
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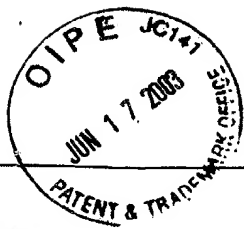
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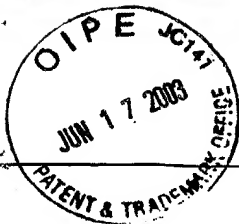
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